



Richardson, T., Shihab, H., Hemani, G., Zheng, J., Hannon, E., Mill, J., Carnero-Montoro, E., Bell, J. T., Lyttleton, O., McArdle, W., Ring, S., Rodriguez, S., Campbell, C., Davey Smith, G., Relton, C., Timpson, N., & Gaunt, T. (2016). Collapsed Methylation Quantitative Trait Loci analysis for Low Frequency and Rare variants. *Human Molecular Genetics*, 25(19), 4339-4349.
<https://doi.org/10.1093/hmg/ddw283>

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Supplementary Tables

Supplementary Table 1: Analysis of Variants within CpG Islands (MAF \leq 5%)

CpG Island	Nearest Gene	Probe	nVars	P-value
chr3:142315171..142315567	<i>PLS1</i>	cg20824294	2	2.86x10 ⁻¹⁹
chr16:88883056..88883461	<i>GALNS</i>	cg00230183	5	3.55x10 ⁻¹⁶
chr16:67427284..67428950	<i>ZDHHC1</i>	cg27539214	5	5.83x10 ⁻¹⁶
chr2:241529962..241530345	<i>CAPN10</i>	cg26250154	2	9.74x10 ⁻¹⁶
chr16:67233032..67233862	<i>ELMO3</i>	cg08721338	2	2.25x10 ⁻¹⁵
chr18:75690494..75690901	<i>BDP1P</i>	cg20484215	3	2.08x10 ⁻¹⁴
chr18:74114551..74114791	<i>ZNF516</i>	cg24786174	3	2.36x10 ⁻¹⁴
chr14:64108343..64108952	<i>WDR89</i>	cg14387360	6	4.12x10 ⁻¹⁴
chr7:36010997..36011407	<i>PP13004</i>	cg08837626	2	5.95x10 ⁻¹⁴
chr17:62339803..62340799	<i>TEX2</i>	cg22194188	2	1.29x10 ⁻¹³

Supplementary Table 2: Analysis of Variants within CpG Islands, Shores & Shelves (MAF \leq 5%)

CpG Island, Shores & Shelves	Nearest Gene	Probe	nVars	P-value
chr1:155260318..155269536	<i>PKLR</i>	cg08949192	12	1.11x10 ⁻²³
chr8:144713866..144722798	<i>ZNF623</i>	cg16316162	4	2.48x10 ⁻²³
chr10:835608..844210	<i>LARP4B</i>	cg10556349	9	3.65x10 ⁻²²
chr21:45726220..45734444	<i>PFKL</i>	cg21069494	8	1.75x10 ⁻²¹
chr16:67139282..67147827	<i>C16orf70</i>	cg00510320	9	9.10x10 ⁻²¹
chr3:53074956..53085101	<i>RFT1</i>	cg04865290	14	1.21x10 ⁻²⁰
chr21:48083200..48092106	<i>PRMT2</i>	cg24877093	13	1.28x10 ⁻²⁰
chr11:47611710..47620055	<i>C1QTNF4</i>	cg04721828	10	1.11x10 ⁻¹⁸
chr2:128346285..128354640	<i>MYO7B</i>	cg01966334	12	1.29x10 ⁻¹⁸
chr2:196517555..196526950	<i>SLC39A10</i>	cg19655195	14	1.66x10 ⁻¹⁸

Supplementary Table 3: Analysis of Variants within CpG Islands (MAF \leq 1%)

CpG Island	Nearest Gene	Probe	nVars	P-value
chr16:3550773..3551274	<i>CLUAP1</i>	cg27559562	3	4.08x10 ⁻¹⁰
chr4:129208297..129209780	<i>PGRMC2</i>	cg13841734	4	4.33x10 ⁻⁰⁹
chr7:148982079..148982675	<i>ZNF783</i>	cg06818605	2	1.05x10 ⁻⁰⁸
chr14:65438412..65439363	<i>FNTB</i>	cg10534904	2	9.25x10 ⁻⁰⁸
chr16:89237944..89238331	<i>CDH15</i>	cg00511475	2	1.61x10 ⁻⁰⁷
chr15:42371634..42371842	<i>PLA2G4D</i>	cg13762474	2	3.48x10 ⁻⁰⁷
chr13:41495776..41496213	<i>TPTE2P5</i>	cg22658229	3	4.02x10 ⁻⁰⁷
chr3:195488595..195489381	<i>MUC4</i>	cg00789671	4	4.11x10 ⁻⁰⁷
chr3:155461917..155462561	<i>PLCH1</i>	cg12679230	2	1.19x10 ⁻⁰⁶
chr8:142401533..142402494	<i>PTP4A3</i>	cg12873476	3	1.56x10 ⁻⁰⁶

Supplementary Table 4: Analysis of Variants within CpG Islands & Shores (MAF \leq 1%)

CpG Island & Shores	Nearest Gene	Probe	nVars	P-value
chr2:233243999..233248448	<i>ALPP</i>	cg16700265	3	3.62x10 ⁻¹⁷
chr16:69758293..69762506	<i>NQO1</i>	cg19194454	4	2.35x10 ⁻¹⁴
chr2:109400907..109405566	<i>CCDC138</i>	cg14300730	7	2.95x10 ⁻¹⁴
chr2:109063509..109067954	<i>GCC2</i>	cg14300730	6	4.44x10 ⁻¹⁴
chr1:1281933..1287345	<i>DVL1</i>	cg02279987	2	6.26x10 ⁻¹⁴
chr1:215738507..215743266	<i>KCTD3</i>	cg12133444	3	1.10x10 ⁻¹³
chr2:109333768..109338670	<i>RANBP2</i>	cg14300730	6	1.96x10 ⁻¹³
chr1:45985163..45989803	<i>PRDX1</i>	cg03335125	9	4.05x10 ⁻¹²
chr22:46260065..46265747	<i>ATXN10</i>	cg12301347	5	4.44x10 ⁻¹²
chr1:46267099..46271820	<i>MAST2</i>	cg03335125	4	9.58x10 ⁻¹²

Supplementary Table 5: Analysis of Variants within CpG Islands, Shores & Shelves (MAF \leq 1%)

CpG Island, Shores & Shelves	Nearest Gene	Probe	nVars	P-value
chr16:69756293..69764506	<i>NQO1</i>	cg19194454	4	2.35x10 ⁻¹⁴
chr1:215736507..215745266	<i>KCTD3</i>	cg12133444	5	2.99x10 ⁻¹³
chr18:76318675..76327015	<i>ATP9B</i>	cg21199045	9	6.19x10 ⁻¹²
chr1:220917411..220926176	<i>C1orf115</i>	cg19030620	9	2.04x10 ⁻¹¹
chr7:87501100..87510493	<i>SLC25A40</i>	cg26365742	7	4.36x10 ⁻¹¹
chr22:40053941..40062844	<i>CACNA1I</i>	cg18436500	4	4.78x10 ⁻¹¹
chr6:5998471..6009125	<i>NRN1</i>	cg09986316	5	8.13x10 ⁻¹¹
chr15:69587246..69595754	<i>PAQR5</i>	cg00642615	5	8.15x10 ⁻¹¹
chr6:43138013..43146217	<i>SRF</i>	cg20227255	2	1.14x10 ⁻¹⁰
chr2:21262669..21270961	<i>APOB</i>	cg07636176	2	1.72x10 ⁻¹⁰

Individual Effects

Supplementary Table 6: Individual effects for CpG Islands results (MAF \leq 5%)

CpG Island	Nearest Gene	Probe	nVars	P-value	Individual effects
chr3:142315171..142315567	<i>PLS1</i>	cg20824294	2	2.86E-19	rs112514146 (3.05E-11), rs192275612 (4.44E-11)
chr16:88883056..88883461	<i>GALNS</i>	cg00230183	5	3.55E-16	rs147207266 (0.04), rs112609821 (9.09E-11), rs117938568 (0.54), rs79554382 (9.09E-11), rs72813559 (4.76E-13)
chr16:67427284..67428950	<i>ZDHHC1</i>	cg27539214	5	5.83x10 ⁻¹⁶	rs114529082 (1.18E-14), rs9922191 (0.45), rs78558968 (1.24E-03), rs145348547 (1.18E-14), rs117132786 (1.48E-03)
chr2:241529962..241530345	<i>CAPN10</i>	cg26250154	2	9.74x10 ⁻¹⁶	rs72999976 (2.21E-05), rs182360139 (7.31E-14)
chr16:67233032..67233862	<i>ELMO3</i>	cg08721338	2	2.25E-15	rs75193761 (1.50E-06), rs8058614 (1.86E-11)
chr18:75690494..75690901	<i>BDPIP</i>	cg20484215	3	2.08E-14	rs11661975 (0.71), rs7506894 (2.21E-14), rs138928621 (3.45E-14)
chr18:74114551..74114791	<i>ZNF516</i>	cg24786174	3	2.36E-14	rs35943460 (1.70E-05), rs117145635 (1.85E-05), rs141520495 (1.64E-08)
chr14:64108343..64108952	<i>WDR89</i>	cg14387360	6	4.12E-14	rs8018194 (6.75E-14), rs8018382 (1.38E-14), rs185458911 (0.99), rs10498511 (1.38E-14), rs61985669 (1.38E-14), rs8018872 (1.38E-14)
chr7:36010997..36011407	<i>PPI3004</i>	cg08837626	2	5.95E-14	rs140384299 (3.99E-14), rs1962367 (1.92E-14)
chr17:62339803..62340799	<i>TEX2</i>	cg22194188	2	1.29E-13	rs9895312 (5.53E-14), rs112866441 (5.53E-14)

Supplementary Table 7: Individual effects for CpG Islands results (MAF \leq 1%)

CpG Island	Nearest Gene	Probe	nVars	P-value	Individual effects
chr16:3550773..3551274	<i>CLUAP1</i>	cg27559562	3	4.08×10^{-10}	rs111497094 (5.88×10^{-04}), rs115697151 (1.59×10^{-09}), rs192397963 (0.31)
chr4:129208297..129209780	<i>PGRMC2</i>	cg13841734	4	4.33×10^{-09}	rs3733261 (4.85×10^{-09}), rs41298551 (4.85×10^{-09}), rs41298547 (0.32), rs41298545 (6.45×10^{-09})
chr7:148982079..148982675	<i>ZNF783</i>	cg06818605	2	1.05×10^{-08}	rs112616806 (4.47×10^{-09}), rs138890106 (0.16)
chr14:65438412..65439363	<i>FNTB</i>	cg10534904	2	9.25×10^{-08}	rs184621947 (0.15), rs188596757 (4.29×10^{-03})
chr16:89237944..89238331	<i>CDH15</i>	cg00511475	2	1.61×10^{-07}	rs190099280 (1.32×10^{-07}), rs117273249 (8.17×10^{-04})
chr15:42371634..42371842	<i>PLA2G4D</i>	cg13762474	2	3.48×10^{-07}	rs75210712 (2.92×10^{-07}), rs112756552 (2.92×10^{-07})
chr13:41495776..41496213	<i>TPTE2P5</i>	cg22658229	3	4.02×10^{-07}	rs77750044 (3.38×10^{-07}), rs111433219 (3.38×10^{-07}), rs74242796 (3.38×10^{-07})
chr3:195488595..195489381	<i>MUC4</i>	cg00789671	4	4.11×10^{-07}	rs115531913 (3.46×10^{-07}), rs116085423 (3.46×10^{-07}), rs76394037 (3.46×10^{-07}), rs147056758 (3.46×10^{-07})
chr3:155461917..155462561	<i>PLCH1</i>	cg12679230	2	1.19×10^{-06}	rs139489498 (0.02), rs58689622 (3.46×10^{-06})
chr8:142401533..142402494	<i>PTP4A3</i>	cg12873476	3	1.56×10^{-06}	rs7830943 (1.48×10^{-06}), rs67017417 (2.34×10^{-06}), rs72660242 (1.48×10^{-06})

Supplementary Table 8: Individual effects for CpG Islands & Shores results (MAF \leq 5%)

CpG Island & Shores	Nearest Gene	Probe	nVars	P-value	Individual effects
chr21:45728220..45732444	<i>PFKL</i>	cg21069494	6	6.24x10 ⁻²⁴	rs150947529 (9.11E-07), rs183612767 (0.82), rs113934313 (7.00E-03), rs140395625 (1.33E-12), rs144344436 (0.07), rs78931138 (1.44E-08)
chr8:144715866..144720798	<i>ZNF623</i>	cg16316162	3	4.02x10 ⁻²³	rs144074829 (1.28E-12), rs149005143 (4.09E-14), rs2384928 (0.33)
chr11:66492937..66498387	<i>SPTBN2</i>	cg24851651	6	2.68x10 ⁻²¹	rs117612543 (0.36), rs479261 (1.25E-10), rs190683929 (0.12), rs138549284 (0.99), rs145618970 (0.02), rs117239013 (2.49E-12)
chr2:75785717..75790312	<i>EVA1A</i>	cg26175789	9	6.73x10 ⁻²¹	rs80132995 (1.30E-08), rs116763000 (0.78), rs114883774 (4.46E-04), rs78222446 (4.46E-04), rs114973170 (8.07E-04), rs76553177 (0.07), rs113609413 (1.17E-12), rs116453138 (4.06E-03), rs72816744 (7.31E-05)
chr12:120753346..120757672	<i>PLA2G1B</i>	cg06379361	5	1.21x10 ⁻²⁰	rs148928469 (1.33E-11), rs141183115 (0.97), rs137933811 (9.48E-11), rs147288206 (0.97), rs140911250 (7.54E-04)
chr7:100873555..100878212	<i>CLDN15</i>	cg01299997	7	1.26x10 ⁻²⁰	rs11773455 (0.96), rs115611906 (7.78E-12), rs183990374 (0.55), rs144733493 (1.28E-09), rs143800908 (0.56), rs80277788 (0.37), rs77731918 (1.50E-03)
chr3:53076956..53083101	<i>RFT1</i>	cg04865290	10	2.03x10 ⁻²⁰	rs183255344 (0.39), rs74380125 (3.83E-13), rs116211523 (0.70), rs142052001 (0.71), rs200910296 (2.10E-09), rs3806693 (0.39), rs189440791 (0.39), rs114006553 (0.14), rs186372583 (0.39), rs180738561 (0.39)
chr2:196519555..196524950	<i>SLC39A10</i>	cg19655195	8	2.23x10 ⁻²⁰	rs181029438 (6.63E-05), rs1371122 (3.01E-12), rs181591378 (0.19), rs116877579 (0.02), rs10184777 (1.71E-11), rs3738940 (0.02), rs78456147 (0.07), rs6719584 (9.54E-13)
chr1:1287707..1293126	<i>MXRA8</i>	cg17132079	3	1.11x10 ⁻¹⁹	rs13249 (3.76E-14), rs147405504 (0.02), rs202220998 (2.54E-07)
chr11:64407877..64413253	<i>NRXN2</i>	cg19395706	4	1.43x10 ⁻¹⁹	rs57387921 (2.12E-14), rs71579882 (6.03E-09), rs71579879 (0.40), rs11820322 (2.12E-14)

Supplementary Table 9: Individual effects for CpG Islands & Shores results (MAF \leq 1%)

CpG Island & Shores	Nearest Gene	Probe	nVars	P-value	Individual effects
chr2:233243999..233248448	<i>ALPP</i>	cg16700265	3	3.62x10 ⁻¹⁷	rs150558405 (4.17E-09), rs73001984 (0.26), rs181094685 (6.21E-11)
chr16:69758293..69762506	<i>NQO1</i>	cg19194454	4	2.35x10 ⁻¹⁴	rs71534273 (0.13), rs117928302 (9.76E-14), rs111555725 (0.13), rs141281805 (0.45)
chr2:109400907..109405566	<i>CCDC138</i>	cg14300730	7	2.95x10 ⁻¹⁴	rs10164715 (0.65), rs58791443 (0.48), rs10195166 (0.48), rs146848524 (1.22E-14), rs139590986 (1.22E-14), rs149132027 (1.22E-14), rs79377757 (1.22E-14)
chr2:109063509..109067954	<i>GCC2</i>	cg14300730	6	4.44x10 ⁻¹⁴	rs191654036 (0.15), rs142672792 (0.54), rs147355200 (6.34E-14), rs76672099 (6.34E-14), rs115218746 (6.34E-14), rs146969797 (6.34E-14)
chr1:1281933..1287345	<i>DVLI</i>	cg02279987	2	6.26x10 ⁻¹⁴	rs145425222 (2.15E-12), rs113488343 (7.45E-09)
chr1:215738507..215743266	<i>KCTD3</i>	cg12133444	3	1.10x10 ⁻¹³	rs76551420 (8.60E-14), rs80018194 (8.60E-14), rs141935194 (0.77)
chr2:109333768..109338670	<i>RANBP2</i>	cg14300730	6	1.96x10 ⁻¹³	rs184224507 (1.22E-14), rs139400907 (0.48), rs113447280 (0.48), rs143297644 (0.48), rs62151247 (0.21), rs59221702 (0.48)
chr1:45985163..45989803	<i>PRDX1</i>	cg03335125	9	4.05x10 ⁻¹²	rs6664588 (0.69), rs6677135 (0.69), rs34181355 (0.69), rs34426720 (0.19), rs148195562 (5.37E-13), rs146962773 (0.69), rs12066809 (0.69), rs12065695 (0.69), rs202109079 (0.69)
chr22:46260065..46265747	<i>ATXN10</i>	cg12301347	5	4.44x10 ⁻¹²	rs136041 (8.72E-06), rs136042 (6.75E-06), rs183318428 (5.12E-12), rs78345649 (5.12E-12), rs80114170 (6.43E-11)
chr1:46267099..46271820	<i>MAST2</i>	cg03335125	4	9.58x10 ⁻¹²	rs76912510 (1.22x10 ⁻⁰⁴), rs151010228 (2.52x10 ⁻¹⁰), rs150406727 (0.54), rs35106493 (0.92)

Supplementary Table 10: Individual effects for CpG Islands, Shores & Shelves results (MAF ≤ 5%)

CpG Island, Shores & Shelves	Nearest Gene	Probe	nVars	P-value	Individual effects
chr1:155260318..155269536	<i>PKLR</i>	cg08949192	12	1.11x10 ⁻²³	rs191588587 (0.29), rs8177992 (0.15), rs8177988 (3.34E-03), rs3762272 (3.84E-05), rs139885057 (0.12), rs8177971 (2.34E-08), rs138344553 (3.03E-09), rs142339464 (3.03E-09), rs115736167 (7.07E-07), rs112966658 (9.43E-03), rs180921974 (5.75E-06), rs186226317 (4.94E-03)
chr8:144713866..144722798	<i>ZNF623</i>	cg16316162	4	2.48x10 ⁻²³	rs144074829 (1.28E-12), rs149005143 (4.09E-14), rs2384928 (0.33), rs4873813 (0.33)
chr10:835608..844210	<i>LARP4B</i>	cg10556349	9	3.65x10 ⁻²²	rs186726285 (2.66E-06), rs114850712 (1.61E-09), rs117535965 (2.03E-04), rs142571536 (7.15E-08), rs117257789 (0.07), rs7916656 (0.17), rs117479624 (1.71E-06), rs116727192 (1.61E-09), rs140695858 (3.57E-04)
chr21:45726220..45734444	<i>PFKL</i>	cg21069494	8	1.75x10 ⁻²¹	rs117689677 (4.36E-03), rs150947529 (9.11E-07), rs183612767 (0.82), rs113934313 (7.00E-03), rs140395625 (1.33E-12), rs144344436 (0.07), rs78931138 (1.44E-08), rs143345133 (6.59E-04)
chr16:67139282..67147827	<i>C16orf70</i>	cg00510320	9	9.10x10 ⁻²¹	rs74932341 (5.34E-10), rs76764655 (9.46E-14), rs185272357 (0.10), rs115599330 (0.30), rs1881466 (1.36E-10), rs8052623 (0.17), rs17618203 (9.46E-14), rs189409766 (0.87), rs140868667 (1.36E-10)
chr3:53074956..53085101	<i>RFT1</i>	cg04865290	14	1.21x10 ⁻²⁰	rs183255344 (0.39), rs74380125 (3.83E-13), rs116211523 (0.70), rs142052001 (0.71), rs200910296 (2.10E-09), rs3806693 (0.39), rs189440791 (0.39), rs114006553 (0.14), rs186372583 (0.39), rs180738561 (0.39), rs138412876 (0.70), rs192671668 (0.39), rs141046534 (0.31), rs141518615 (0.39)

chr21:48083200..48092106	<i>PRMT2</i>	cg24877093	13	1.28x10 ⁻²⁰	rs76187868 (1.06E-03), rs149520727 (0.66), rs117282473 (1.89E-03), rs79726879 (0.42), rs117164191 (3.06E-03), rs117073279 (0.19), rs145906853 (8.59E-04), rs9306158 (1.26E-03), rs2839379 (2.03E-11), rs187437953 (2.03E-11), rs59497292 (5.88E-04), rs141423093 (0.32), rs112771868 (4.60E-08)
chr11:47611710..47620055	<i>CIQTNF4</i>	cg04721828	10	1.11x10 ⁻¹⁸	rs150142619 (5.99E-03), rs11601694 (1.15E-11), rs112190302 (0.69), rs149860042 (0.14), rs118102336 (8.71E-10), rs149175346 (9.56E-11), rs151202997 (0.28), rs73463785 (0.86), rs111955624 (0.69), rs4752852 (0.69)
chr2:128346285..128354640	<i>MYO7B</i>	cg01966334	12	1.29x10 ⁻¹⁸	rs114134373 (0.84), rs114950198 (0.03), rs183049678 (4.11E-09), rs4662584 (0.01), rs115500899 (4.11E-09), rs79437920 (0.49), rs72843363 (4.05E-09), rs111804739 (2.53E-04), rs59486761 (2.25E-09), rs17340406 (5.21E- 09), rs72843365 (9.24E-03), rs112316764 (0.65)
chr2:196517555..196526950	<i>SLC39A10</i>	cg19655195	14	1.66x10 ⁻¹⁸	rs58650718 (0.02), rs143105938 (8.15E-05), rs184788263 (0.60), rs141335019 (1.71E-11), rs181029438 (6.63E-05), rs1371122 (3.01E-12), rs181591378 (0.19), rs116877579 (0.02), rs10184777 (1.71E-11), rs3738940 (0.02), rs78456147 (0.07), rs6719584 (9.54E-13), rs13489 (4.42E-06), rs10171083 (1.71E-11)

Supplementary Table 11: Individual effects for CpG Islands, Shores & Shelves results (MAF ≤ 1%)

CpG Island, Shores & Shelves	Nearest Gene	Probe	nVars	P-value	Individual effects
chr16:69756293..69764506	<i>NQO1</i>	cg19194454	4	2.35E-14	rs190237675 (0.51), rs71534273 (0.13), rs117928302 (9.76E-14), rs111555725 (0.13)
chr1:215736507..215745266	<i>KCTD3</i>	cg12133444	5	2.99E-13	rs193052948 (4.86E-12), rs76551420 (8.60E-14), rs80018194 (8.60E-14), rs141935194 (0.77), rs59026507 (8.60E-14)
chr18:76318675..76327015	<i>ATP9B</i>	cg21199045	9	6.19E-12	rs117161401 (0.19), rs117600463 (0.55), rs12954375 (0.82), rs28720977 (0.89), rs73972232 (1.07E-09), rs114357617 (0.97), rs80324891 (0.57), rs77501012 (0.55), rs73972233 (7.16E-13)
chr1:220917411..220926176	<i>Clorf115</i>	cg19030620	9	2.04E-11	rs72472366 (1.17E-11), rs78814532 (1.17E-11), rs74487083 (1.17E-11), rs12023149 (1.17E-11), rs2172700 (0.73), rs75917278 (1.17E-11), rs80115160 (1.17E-11), rs2172699 (0.73), rs114783097 (0.06)
chr7:87501100..87510493	<i>SLC25A40</i>	cg26365742	7	4.36E-11	rs117188385 (0.16), rs10280397 (0.71), rs79810525 (3.23E-12), rs117953527 (0.28), rs150166460 (0.87), rs142344486 (0.32), rs111815979 (0.32)
chr22:40053941..40062844	<i>CACNA1I</i>	cg18436500	4	4.78E-11	rs57299573 (8.24E-14), rs58021347 (0.66), rs117956704 (0.61), rs139876154 (0.77)
chr6:5998471..6009125	<i>NRN1</i>	cg09986316	5	8.13x10 ⁻¹¹	rs140163412 (2.22x10 ⁻⁰⁵), rs184320557 (0.26), rs114224069 (3.80x10 ⁻⁰³), rs79453941 (0.02), rs114396677 (1.39x10 ⁻¹¹)
chr15:69587246..69595754	<i>PAQR5</i>	cg00642615	5	8.15x10 ⁻¹¹	rs73437176 (0.08), rs76909122 (0.08), rs111668944 (7.76x10 ⁻¹¹), rs4776438 (0.06), rs117586126 (0.24)
chr6:43138013..43146217	<i>SRF</i>	cg20227255	2	1.14x10 ⁻¹⁰	rs192971989 (1.02x10 ⁻¹⁰), rs112496612 (0.08)
chr2:21262669..21270961	<i>APOB</i>	cg07636176	2	1.72x10 ⁻¹⁰	rs72653053 (0.15), rs1800480 (1.28x10 ⁻¹⁰)

Trans-effects

Supplementary Table 12: Trans-effects for CpG Islands & Shores (MAF \leq 5%)

CpG Islands & Shores	Nearest Gene	Probe	Probe location	Nearest Gene to Probe	nVars	P-value
chr21:33982367..33987450	<i>C21orf59</i>	cg09050820	chr6:167586206	<i>TCP10L2</i>	9	1.50x10 ⁻¹⁵
chr11:47613710..47618055	<i>CIQTNF4</i>	cg17212184	chr11:89787638	<i>TRIM49C</i>	7	8.98x10 ⁻¹⁴
chr11:47609138..47614438	<i>CIQTNF4</i>	cg13118410	chr11:49839281	<i>OR4C13</i>	7	2.53x10 ⁻¹³
chr1:174990365..174994796	<i>MRPS14</i>	cg00981945	chr1:173977793	<i>RC3H1</i>	2	1.20x10 ⁻¹²
chr3:49392855..49397942	<i>RHOA</i>	cg09665691	chr3:47563885	<i>SCAP</i>	3	2.89x10 ⁻¹²
chr11:46408921..46416687	<i>CHRM4</i>	cg09580214	chr11:47448534	<i>PSMC3</i>	5	4.54x10 ⁻¹²
chr11:47268234..47272633	<i>NR1H3</i>	cg04721828	chr11:48285200	<i>OR4X1</i>	5	4.56x10 ⁻¹²
chr11:57100841..57105430	<i>SSRP1</i>	cg26831825	chr11:55541269	<i>OR5D13</i>	6	5.13x10 ⁻¹²
chr1:173443978..173449105	<i>PRDX6</i>	cg02482603	chr1:174843754	<i>RABGAP1L</i>	12	8.14x10 ⁻¹²
chr6:31800598..31804823	<i>C6orf48</i>	cg10154826	chr6:17600994	<i>FAM8A1</i>	18	8.18x10 ⁻¹²

Supplementary Table 13: Trans-effects for CpG Islands & Shores (MAF \leq 1%)

CpG Islands & Shores	Nearest Gene	Probe	Probe location	Nearest Gene to Probe	nVars	P-value
chr6:71996106..72001109	<i>OGFRL1</i>	cg05891823	chr19:9938431	<i>UBL5</i>	7	2.95x10 ⁻¹⁴
chr2:179313214..179318573	<i>PRKRA</i>	cg05098815	chrY:9449618	<i>RBMV3AP</i>	7	5.49x10 ⁻¹⁴
chr22:46608124..46612789	<i>PPARA</i>	cg03601053	chrY:15016101	<i>DDX3Y</i>	2	6.26x10 ⁻¹⁴
chr19:9783480..9787806	<i>ZNF562</i>	cg02900749	chr2:68251473	<i>NA</i>	3	1.32x10 ⁻¹³
chr3:47321747..47326607	<i>KIF9</i>	cg06360465	chr3:50341183	<i>HYAL1</i>	3	3.01x10 ⁻¹³
chr3:48280425..48284817	<i>ZNF589</i>	cg26928789	chrY:22682393	<i>TTYTY10</i>	4	5.82x10 ⁻¹²
chr6:71996106..72001109	<i>OGFRL1</i>	cg05891823	chr19:9938431	<i>UBL5</i>	2	6.47x10 ⁻¹²
chr1:228360442..228364691	<i>IBA57</i>	cg12878555	chr12:6862401	<i>MLF2</i>	6	9.78x10 ⁻¹²
chr20:60811371..60816247	<i>OSBPL2</i>	cg13062406	chr11:18813544	<i>PTPN5</i>	5	2.13x10 ⁻¹¹
chr19:47613253..47619101	<i>ZC3H4</i>	cg13899293	chr4:79175365	<i>FRAS1</i>	6	2.29x10 ⁻¹¹

Evaluations across time points

Supplementary Table 14: Evaluation of Cis-mQTLs using other time points in ARIES

			Birth (N=771)		Adolescence (N=837)		Pregnancy (N=764)		Middle Age (N=742)	
Region	Nearest Gene	Probe	nVars	P-value	nVars	P-value	nVars	P-value	nVars	P-value
CpG Island – MAF 5%										
chr3:142315171..142315567	<i>PLS1</i>	cg20824294	2	3.87E-08	2	2.92E-15	2	7.40E-15	2	6.74E-14
chr16:88883056..88883461	<i>GALNS</i>	cg00230183	5	4.17E-05	5	4.36E-09	5	1.77E-07	5	1.68E-07
chr16:67427284..67428950	<i>ZDHHC1</i>	cg27539214	5	1.34E-15	5	5.63E-13	5	2.71E-08	5	3.75E-12
chr2:241529962..241530345	<i>CAPN10</i>	cg26250154	2	3.76E-12	2	1.28E-11	2	3.06E-16	2	1.31E-14
chr16:67233032..67233862	<i>ELMO3</i>	cg08721338	2	1.52E-10	2	3.21E-16	2	1.79E-10	2	4.78E-07
CpG Island & Shores – MAF 5% (Top 10 Results only (out of 90))										
chr21:45728220..45732444	<i>PFKL</i>	cg21069494	6	9.34E-20	6	1.63E-18	6	2.15E-18	6	2.39E-20
chr8:144715866..144720798	<i>ZNF623</i>	cg16316162	3	5.72E-22	3	2.62E-19	3	1.29E-23	3	3.73E-28
chr11:66492937..66498387	<i>SPTBN2</i>	cg24851651	6	1.06E-12	6	7.04E-17	6	4.52E-12	6	1.76E-11
chr2:75785717..75790312	<i>EVA1A</i>	cg26175789	8	3.27E-14	8	1.08E-20	9	7.24E-19	9	5.47E-18
chr12:120753346..120757672	<i>PLA2G1B</i>	cg06379361	5	3.99E-20	5	8.86E-16	5	3.66E-19	5	6.11E-21
chr7:100873555..100878212	<i>CLDN15</i>	cg01299997	7	4.08E-22	7	3.75E-15	6	1.96E-12	6	3.33E-19
chr3:53076956..53083101	<i>RFT1</i>	cg04865290	10	1.34E-12	10	6.69E-24	10	3.82E-22	10	1.99E-13

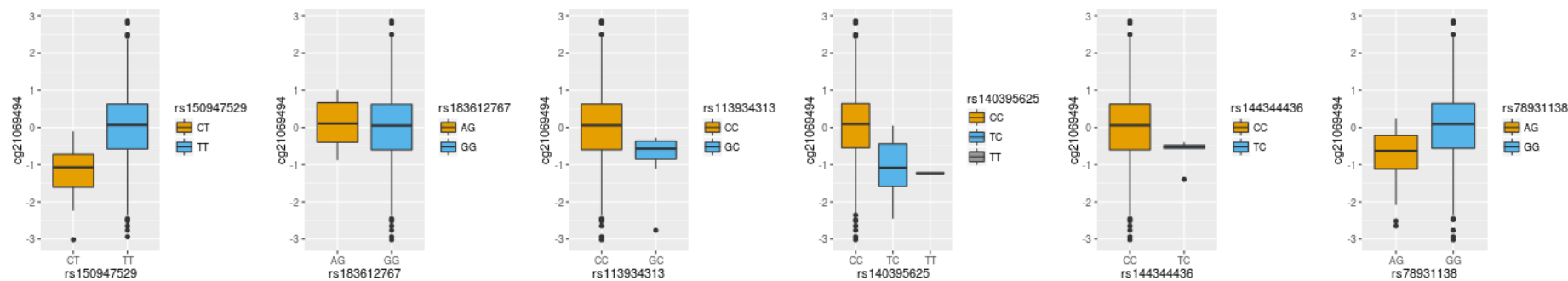
chr2:196519555..196524950	<i>SLC39A10</i>	cg19655195	8	1.25E-14	8	1.68E-18	8	1.84E-15	8	6.86E-16
chr1:1287707..1293126	<i>MXRA8</i>	cg17132079	3	2.99E-13	3	4.02E-20	3	1.06E-14	3	4.54E-14
chr11:64407877..64413253	<i>NRXN2</i>	cg19395706	4	7.53E-14	4	3.00E-16	3	5.63E-17	3	7.56E-23
CpG Island & Shores – MAF 1%										
chr2:233243999..233248448	<i>ALPP</i>	cg16700265	3	3.09x10 ⁻⁰⁹	3	3.08x10 ⁻¹⁴	3	1.08x10 ⁻¹¹	3	7.41x10 ⁻¹⁵
CpG Island, Shores & Shelves – MAF 5% (Top 10 Results only (out of 37))										
chr1:155260318..155269536	<i>PKLR</i>	cg08949192	12	1.43E-12	12	3.04E-18	12	3.26E-27	12	1.90E-21
chr8:144713866..144722798	<i>ZNF623</i>	cg16316162	4	3.10E-22	4	2.30E-19	5	4.06E-24	5	3.55E-28
chr10:835608..844210	<i>LARP4B</i>	cg10556349	9	9.82E-18	9	1.23E-21	9	1.86E-20	9	1.57E-18
chr21:45726220..45734444	<i>PFKL</i>	cg21069494	8	8.83E-17	8	5.10E-16	8	1.13E-18	8	2.86E-20
chr16:67139282..67147827	<i>C16orf70</i>	cg00510320	9	1.39E-22	9	2.37E-21	10	4.45E-18	10	9.94E-19
chr3:53074956..53085101	<i>RFT1</i>	cg04865290	14	1.36E-12	14	1.85E-24	14	2.10E-22	14	2.18E-13
chr21:48083200..48092106	<i>PRMT2</i>	cg24877093	13	8.83E-17	13	6.64E-20	13	3.73E-14	13	3.52E-14
chr11:47611710..47620055	<i>CIQTNF4</i>	cg04721828	10	1.02E-11	10	2.27E-16	10	1.54E-13	10	3.59E-11
chr2:128346285..128354640	<i>MYO7B</i>	cg01966334	12	3.13E-07	12	1.44E-20	12	4.86E-19	12	1.23E-15
chr2:196517555..196526950	<i>SLC39A10</i>	cg19655195	14	4.80E-13	14	3.64E-18	14	4.20E-15	14	9.42E-15

Supplementary Table 15: Evaluation of Trans-mQTLs using other time points in ARIES

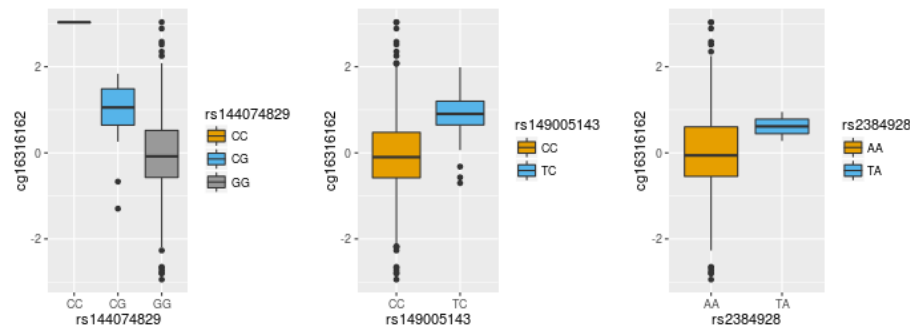
			Birth (N=771)		Adolescence (N=837)		Pregnancy (N=764)		Middle Age (N=742)	
Region	Nearest Gene	Probe	nVars	P-value	nVars	P-value	nVars	P-value	nVars	P-value
CpG Island & Shore – MAF 5%										
chr21:33982367..33987450	<i>C21orf59</i>	cg09050820	9	3.67x10 ⁻¹⁵	9	1.14x10 ⁻¹⁵	9	1.54x10 ⁻¹⁰	9	4.32x10 ⁻¹⁴

Supplementary Figure 1: Box & Whisker plots for CpG Island & Shores cis-effects

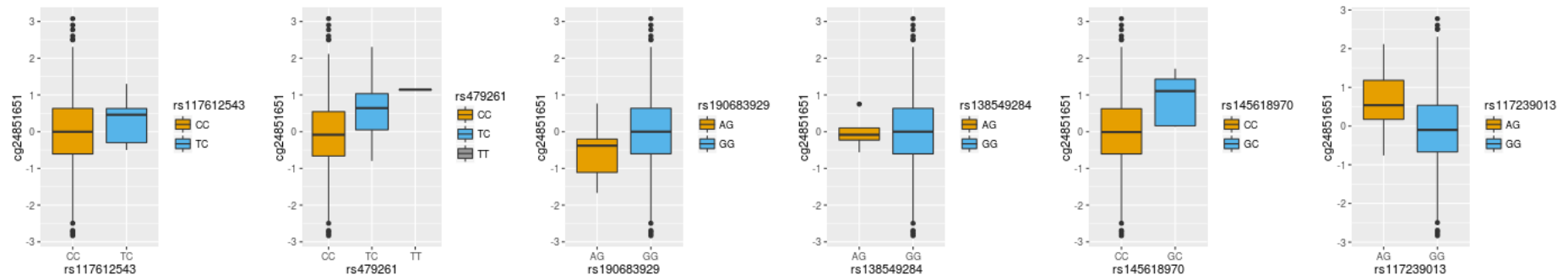
a) *PFKL*



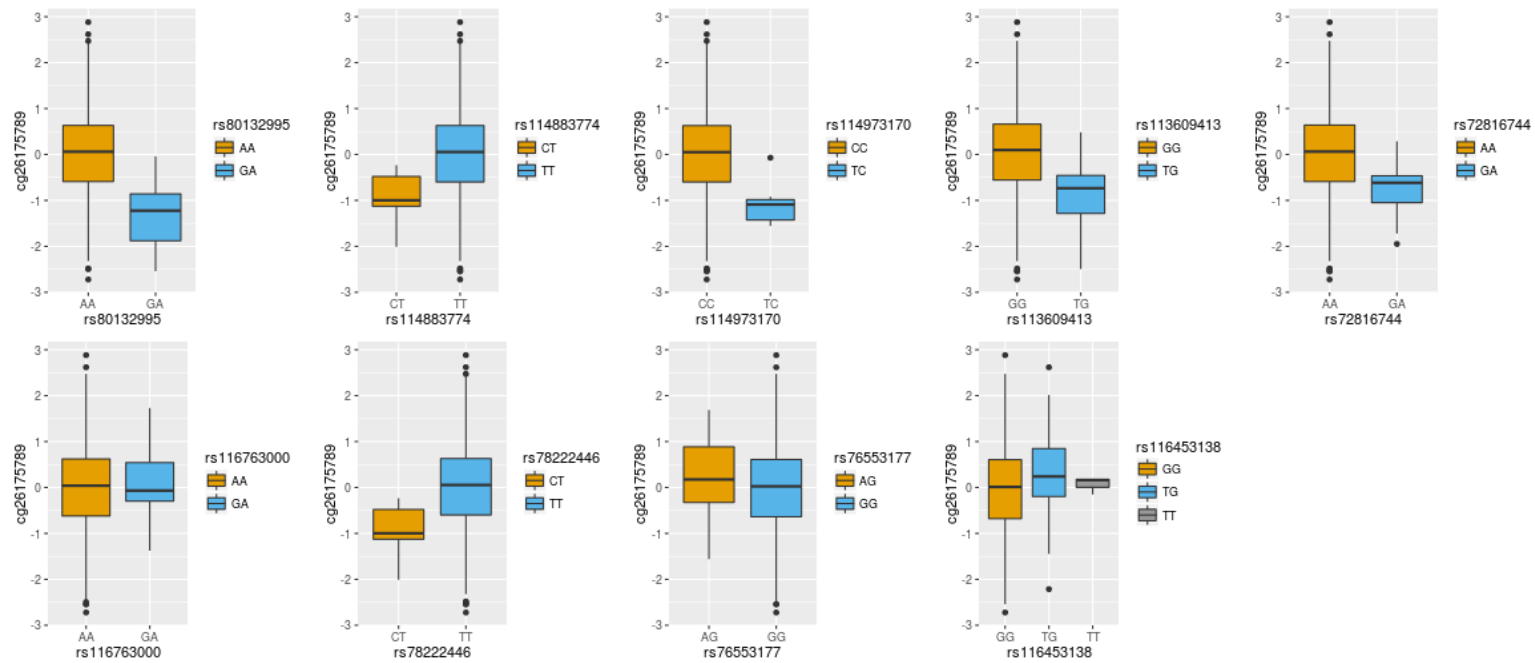
b) *ZNF623*



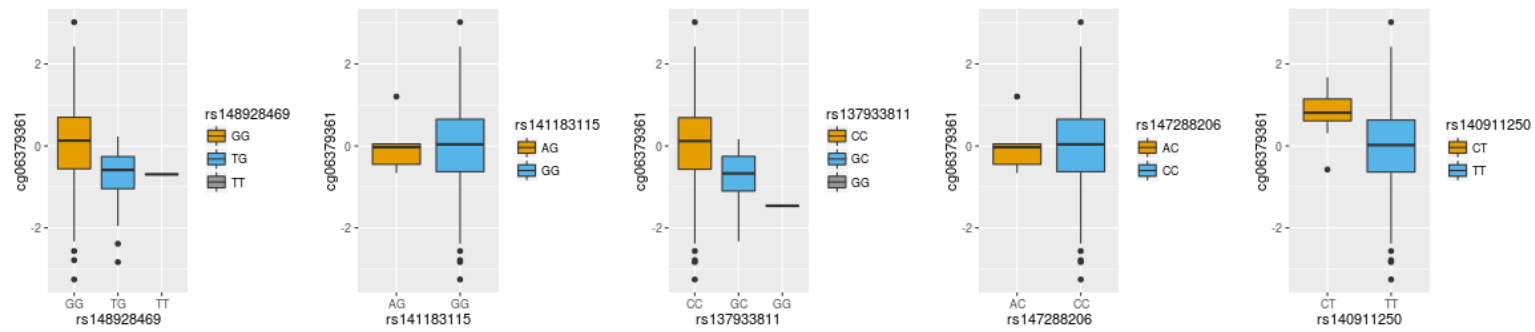
c) *SPTBN2*



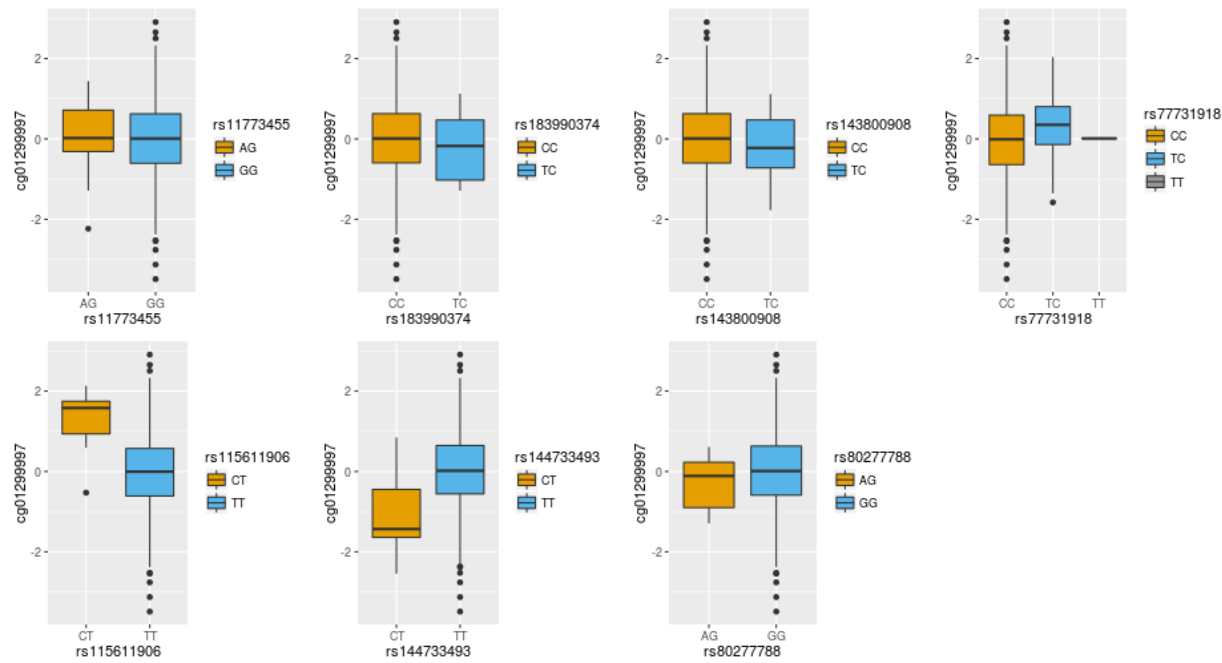
d) *EVA1A*



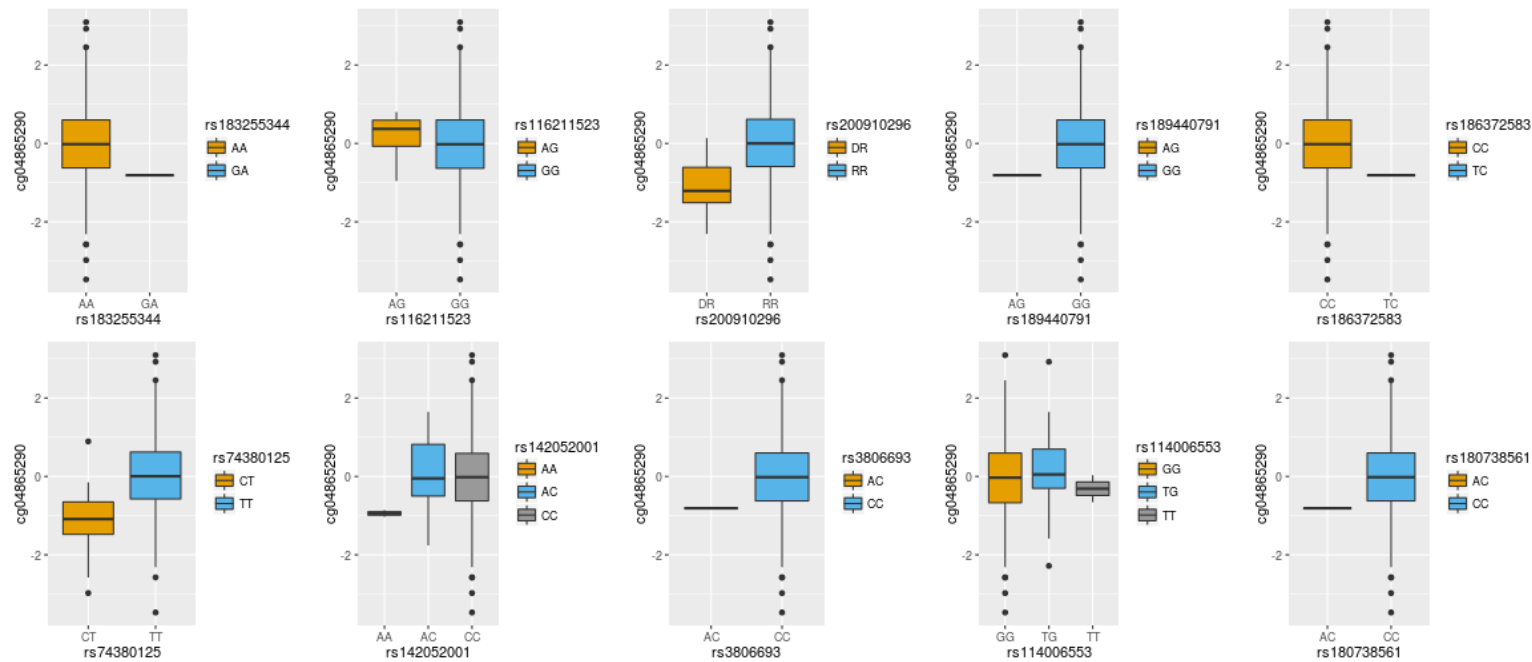
e) *PLA2G1B*



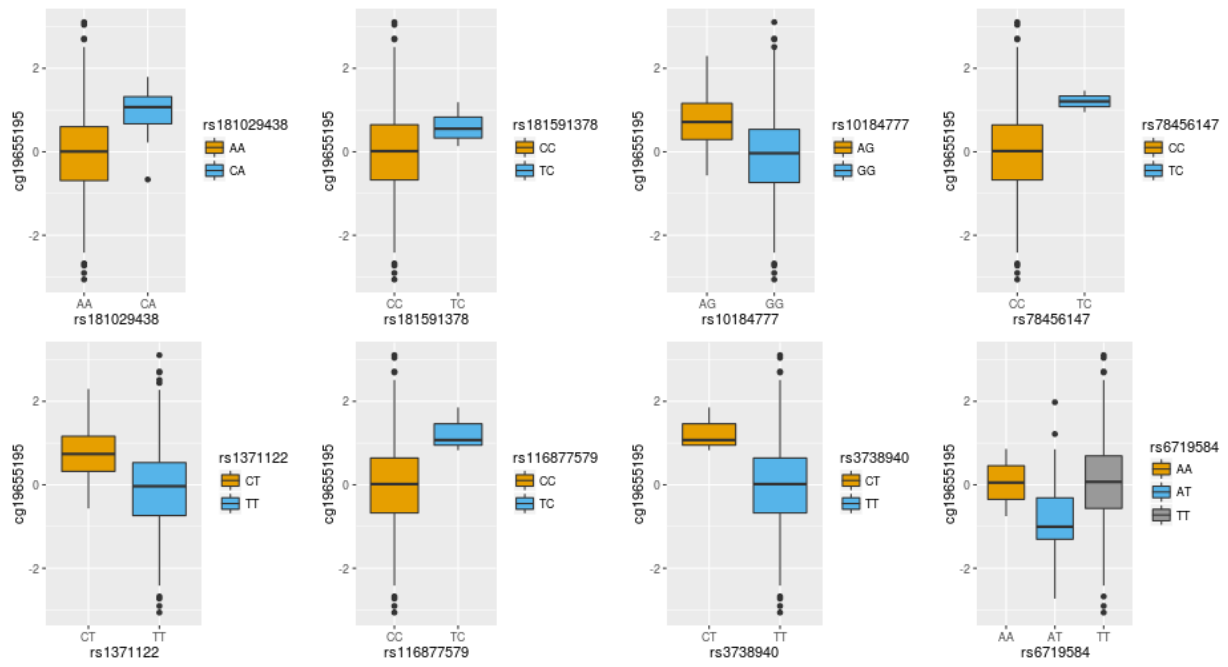
f) *CLDN15*



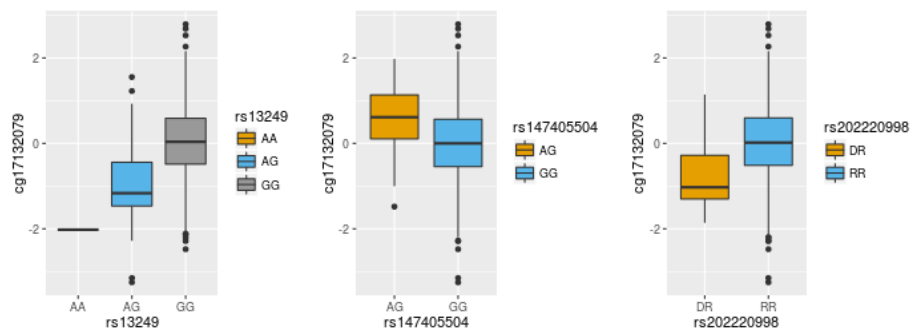
g) *RFT1*



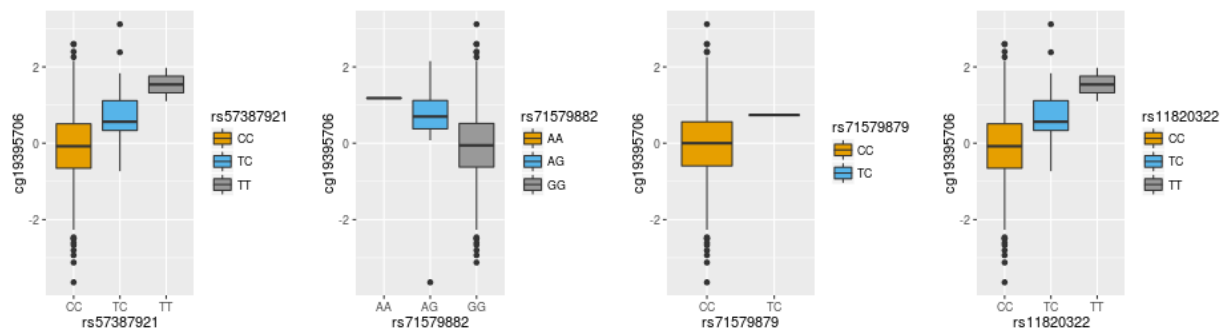
h) *SLC39A10*



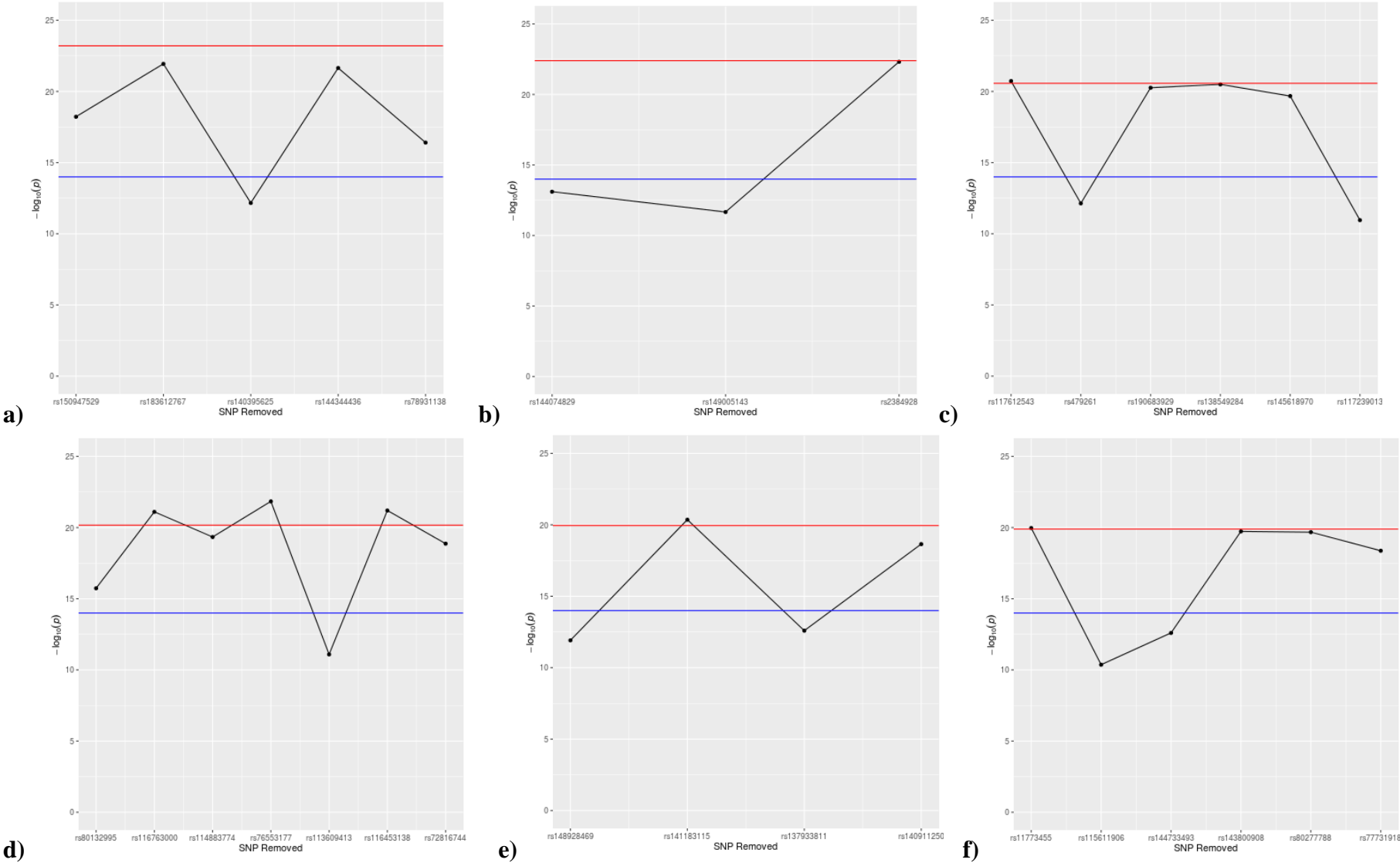
i) *MXRA8*

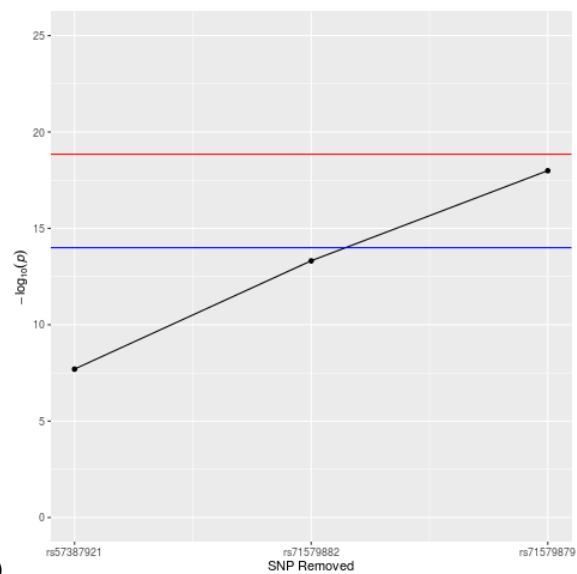
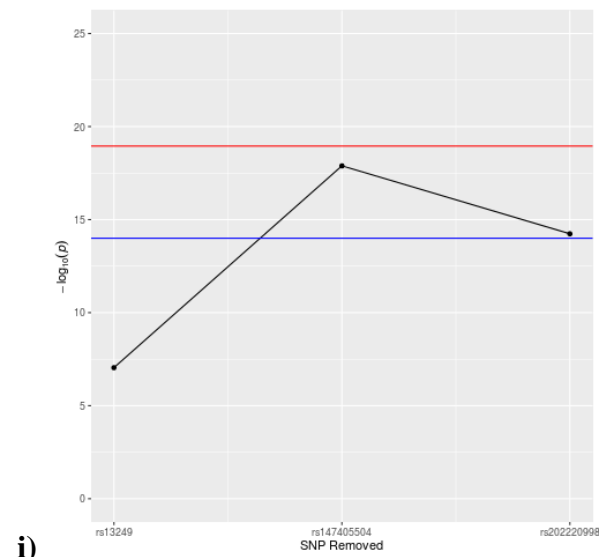
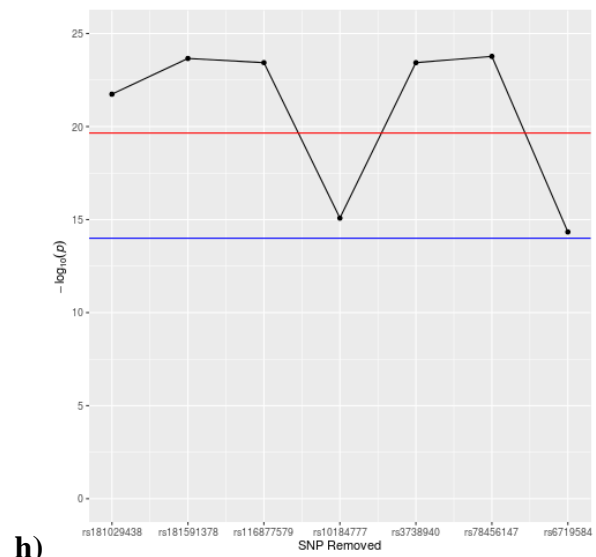
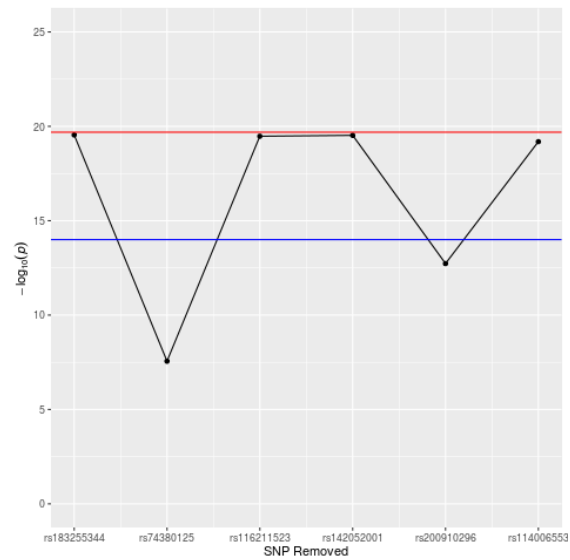


j) *NRXN2*



Supplementary Figure 2: Leave-One-Out Analysis for CpG Island & Shores cis-effects





- Red line = observed p-value according to SKAT when analysing all variants at this mQTL (prior to pruning)
- Blue line = threshold used in our study (i.e. 1.0×10^{-14})
- Each plot resembles one of the top 10 hits from our study at the following loci a) *PFKL* b) *ZNF623* c) *SPTBN2* d) *EVA1A* e) *PLA2G1B* f) *CLDN15* g) *RFT1* h) *SLC39A10* i) *MXRA8* j) *NRXN2*

Supplementary Table 16: Results from Conditional analysis at previously discovered mQTL loci

Conditional analysis results						Single Variant mQTL results		
Region	Nearest Gene	Probe	nVars05	P-value05	Timepoints_collapse	SNP	P-value	Timepoints_single
chr2:208974900..208979396	<i>C2orf80</i>	cg10392614	5	2.56x10 ⁻²⁰	ALL	rs28575061	1.36x10 ⁻¹⁶⁹	ALL
chr20:37350130..37359372	<i>SLC32A1</i>	cg15490840	5	2.88x10 ⁻²⁰	F7	rs10932241	1.64x10 ⁻¹⁷	15up
chr8:26045804..26050097	<i>PPP2R2A</i>	cg12285565	3	1.16x10 ⁻¹⁸	ALL	rs2867326	1.00x10 ⁻¹⁹	cord, F7, 15up, ante
chr16:1003902..1008281	<i>LMF1</i>	cg07338658	5	3.34x10 ⁻¹³	-	rs111820009	1.06x10 ⁻³³	ALL
chr19:41302467..41307050	<i>RAB4B</i>	cg11298343	5	6.32x10 ⁻¹²	-	rs111833532	9.61x10 ⁻⁵⁰	ALL
chr3:13321438..13326929	<i>NUP210</i>	cg05265484	3	8.54x10 ⁻¹²	-	rs36024363	1.77x10 ⁻⁴³	ALL
chr2:1799618..1804060	<i>MYT1L</i>	cg04722030	9	1.17x10 ⁻¹¹	-	rs13387965	1.43x10 ⁻²⁰	F7, 15up
chr13:111299316..111303593	<i>CARS2</i>	cg15747390	18	1.22x10 ⁻¹¹	-	rs61970542	5.87x10 ⁻⁹⁷	ALL
chr3:112928437..112933506	<i>BOC</i>	cg23260991	3	1.56x10 ⁻¹¹	-	rs931702	7.48x10 ⁻¹⁶	FOM
chr1:91187139..91191400	<i>BARHL2</i>	cg22507154	4	1.42x10 ⁻¹⁰	-	rs72720396	8.12x10 ⁻²⁵	F7, 15up, ante, FOM

nVars05 = number of variants analysed (MAF≤5%), P-value05 = SKAT p-value conditioned on single variant mQTL at this loci (MAF≤5%), SNP = mQTL variant at this loci associated with methylation from probe in single variant analysis, P-value = single variant p-value between SNP and probe from single variant analysis, Timepoints collapse = Time points where results from the conditional analyses had a p-value < 1 x 10⁻¹⁴, Timepoints_single = Time points where results from single variant analysis had a p-value < 1 x 10⁻¹⁴. Where multiple time points are reported, the p-value from the time points with the largest sample size is presented.